# (19) World Intellectual Property Organization

International Bureau



(43) International Publication Date 24 June 2004 (24.06.2004)

### (10) International Publication Number WO 2004/052928 A3

- C07K 14/71, (51) International Patent Classification<sup>7</sup>: C12N 15/12, 5/10, C12Q 1/68, G01N 33/53, A61K 38/18, 48/00
- (21) International Application Number:

PCT/GB2003/005374

(22) International Filing Date:

10 December 2003 (10.12.2003)

(25) Filing Language:

(30) Priority Data:

0228776.1

English

(26) Publication Language:

English

10 December 2002 (10.12.2002)

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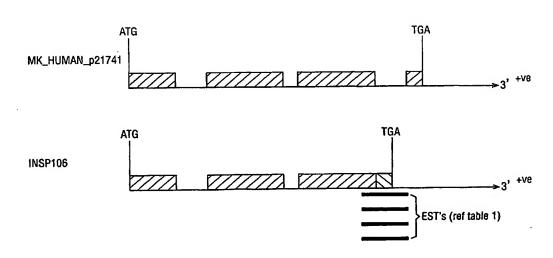
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- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

#### Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- (88) Date of publication of the international search report: 5 August 2004

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: MIDKINE-LIKE PROTEIN



WO 2004/052928 A3 |||||||||||| (57) Abstract: The invention is based on the discovery that the INSP106 protein is a novel splice variant of a known midkine family member (swalliP21741IMK\_HUMAN).

#### INTERNATIONAL SEARCH REPORT

ional Application No PCT/GB 03/05374

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C07K14/71 C12N15/12 A61K38/18 A61K48/00

C. DOCUMENTS CONSIDERED TO BE RELEVANT

C12N5/10

C12Q1/68

G01N33/53

According to international Patent Classification (IPC) or to both national classification and IPC

#### B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)  $IPC \ 7 \ C07K \ C12N \ C12Q \ A61K \ G01N$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, EMBASE, EMBL, GENSEQ, WPI Data, PAJ

Category °	Citation of document, with Indication, where appropriate, of the	e relevant passages	Relevant to claim No.		
Х	DATABASE UNIPROT 'Online! 1 October 2000 (2000-10-01) IKEDA,S. ET AL.: "Midkine prec Database accession no. Q9N0E6 XP002282467	ursor"	1,2,4-45		
<b>X</b> .	EP 0 476 233 A (AMERICAN CYANA 25 March 1992 (1992-03-25) claims 1-13	MID CO)	4,7-15, 19-45		
X	US 6 083 907 A (MURAMATSU TAKA 4 July 2000 (2000-07-04) SEQ ID NO:2 and SEQ ID NO:3 column 4, line 10 -column 8, l	·	4,7-15, 19-45		
χ Furt	her documents are listed in the continuation of box C.	χ Patent family members are listed	d in annex.		
<ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the art which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the international filing date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filing date but later than the priority date claimed</li> </ul>		cited to understand the principle or to invention  "X" document of particular relevance; the cannot be considered novel or cannot have an inventive step when the cannot be considered to involve an invention to considered to involve an indocument is combined with one or respectively.	t of particular relevance; the claimed invention be considered novel or cannot be considered to an inventive step when the document is taken alone to f particular relevance; the claimed invention be considered to involve an inventive step when the nit is combined with one or more other such docusuch combination being obvious to a person skilled to member of the same patent family		
	actual completion of the international search  7 May 2004	14/06/2004	аын төрип		
	malling address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer  ALCONADA RODRIG.	., A		

### INTERNATIONAL SEARCH REPORT

In onal Application No PCT/GB 03/05374

		PCT/GB 03/05374					
C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT							
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.					
X	WO 01/36635 A (BURGESS CATHERINE ;PRAYAGA SUDHIRDAS K (US); CURAGEN CORP (US); MA) 25 May 2001 (2001-05-25) page 3, last paragraph -page 8, paragraph 3 page 48, paragraph 2 -page 49, paragraph 1 page 50, paragraph 2 -page 83, paragraph 2 page 84, paragraph 1 -page 111, paragraph 2	4,7-15, 19-45					
Α	AKHTER SHAMIMA ET AL: "Clusters of basic amino acids in midkine: Roles in neurite-promoting activity and plasminogen activator-enhancing activity" JOURNAL OF BIOCHEMISTRY (TOKYO), vol. 123, no. 6, June 1998 (1998-06), pages 1127-1136, XP001189525 ISSN: 0021-924X cited in the application						
A	BERNARD-PIERROT ISABELLE ET AL: "Dominant negative effectors of heparin affin regulatory peptide (HARP) angiogenic and transforming activities" JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD, US, vol. 277, no. 35, 30 August 2002 (2002-08-30), pages 32071-32077, XP002245580 ISSN: 0021-9258 cited in the application						

# mational application No. PCT/GB 03/05374

### INTERNATIONAL SEARCH REPORT

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)					
This international Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:						
1. χ	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:					
	Although claims 35-37 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.					
2. X	Claims Nos.: 16-18 (complete) and 14, 19, 31, 33-37 (partially) because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:					
	see FURTHER INFORMATION sheet PCT/ISA/210					
з. 🗌	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).					
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)					
This inte	emational Searching Authority found multiple inventions in this international application, as follows:					
	• •					
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.					
2	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.					
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:					
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:					
Remari	k on Protest  The additional search fees were accompanied by the applicant's protest.					
	No protest accompanied the payment of additional search fees.					

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 16-18 (complete) and 14, 19, 31, 33-37 (partially)

Present claim 14, 19, 31, 33-37 relate to a ligand and uses thereof which is defined by reference to a desirable characteristic or property, namely, that binds specifically to a polypeptide according to any one of claims 1-7. The claims cover all compounds having this characteristic or property, whereas the application provides support within the meaning of Article 6 PCT and disclosure within the meaning of Article 5 PCT for only a very limited number of such compounds. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the compound by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible. Consequently, the search has been carried out for those of the claims as far as they relate to the ligands which are supported and disclosed, i.e the antibodies specific for the polypeptide of claims 1-7 (e.g claim 15).

Present claims 16-19, 31, 33-37 relate to a compound and uses thereof which is defined by reference to a desirable characteristic or property, namely, that either increases, decreases the level of expression or activity of the polypeptides according to claims 1-7. The claims cover all compounds having this characteristic or property, whereas the application provides support within the meaning of Article 6 PCT and disclosure within the meaning of Article 5 PCT for only a very limited number of such compounds. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the compound by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible. Consequently, the search has not been carried out for those of the claims as far as they relate to the unidentified compound.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

## INTERNATIONAL SEARCH REPORT

tn onal Application No
PCT/GB 03/05374

Patent document cited in search report	Publication date		Patent family member(s)	Publication date
EP 0476233 A	25-03-1992	US	5210026 A	11-05-1993
2. 0.,0200		AT	134219 T	15-02-1996
		AU	636789 B2	06-05-1993
		AU	8258591 A	27-02-1992
		CA	2049370 A1	21-02-1992
		DE	69117123 D1	28-03-1996
		DE	69117123 T2	02-10-1996
		DK	476233 T3	11-03-1996
		EP	0476233 A1	25-03-1992
		ES	2085928 T3	16-06-1996
		GR	3019059 T3	31-05-1996
		JP	6217778 A	09-08-1994
		NZ	239357 A	27-07-1993
		PT	98694 A	31-07-1992
		ZA	9106552 A	27-05-1992
US 6083907	04-07-2000	CA	2220854 A1	26-07-1999
		JP	9095454 A	08-04-1997
WO 0136635	25-05-2001	AU	1486801 A	30-05-2001
310000		WO	0136635 A2	25-05-2001